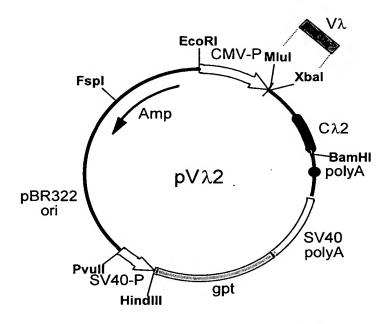


**FIG.** 1



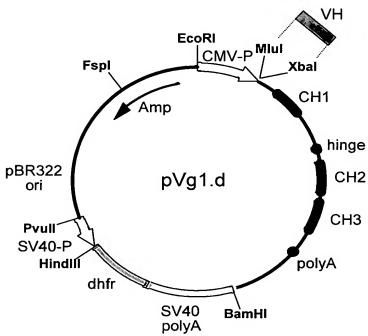


FIG. 2

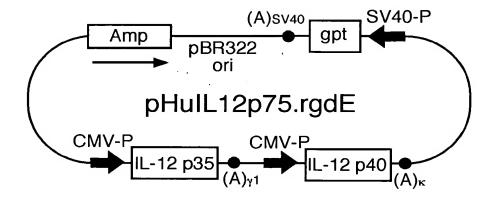


FIG. 3

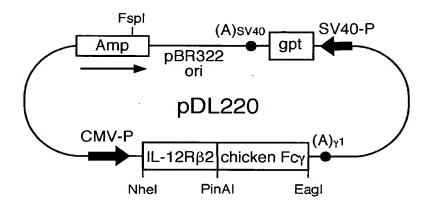


FIG. 4

## (A) Vλ sequences

		1 0123456789	2 01234567890	3 1234567899	4 0123456789
Chimeric Bl Humanized Bl DPL16	**ALTQPAS SSELTQ <u>P</u> P <u>S</u>	*VSANLGGTV *VSVALGQTV	KITCSGGYSG*	A *YYG WYQQK: *YYGWYQQK	S PGSAPVTVIY PGQAPVT VIY
	5	6	7	8	9 01234556789 A
Chimeric B1 Humanized B1 DPL16	DNTRRPSDIP	SRFSGSKSG	S TATLTITGY	AEDEADYYCG	TWDSSRVGI FG TWDSSRVGIFG
Chimeric B1 Humanized B1 DPL16/Jλ2					
(B) VH sequences					
Chimeric B1 Humanized B1 DP-54	123456789 AVTLDESGG EVQLVESGG	GLQTPGGALS GLVQPGGSLR	0123456789 LVCKASGFTF	0123456789 ( SSYSML WVRQ SSYSMLWVRQ	APGKGLEYVA APGKGLEY_VA
Chimeric B1 YYCARSSVYS Humanized B1 YYCARSSVYS DP-54	A EITNTGRTRR	Y GAAVKG RA Y GAAVKGRA	7 39 012345678 TI SRDNGQSTV TI SRDNAKN <u>T</u> T	ABC VR LQLNNLRAE VY LQMNSLRAE	DTGT
Chimeric B1 Humanized B1 JH1	ABCDEFGHI CSYGWCAGNII CSYGWCAGNII		1 0123 IVSS IVSS		

FIG. 5

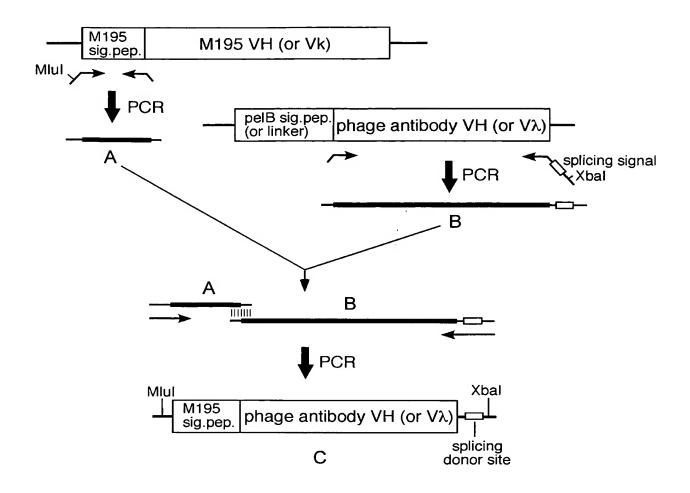


FIG. 6

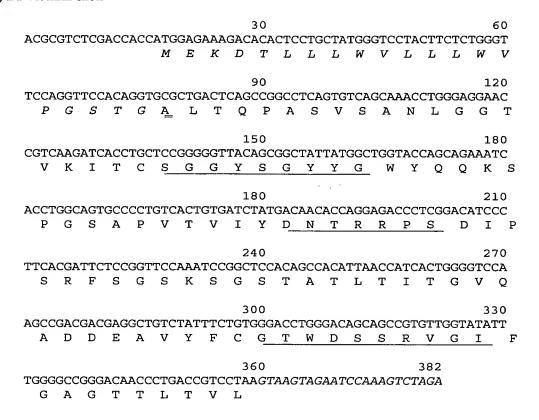


FIG. 7A

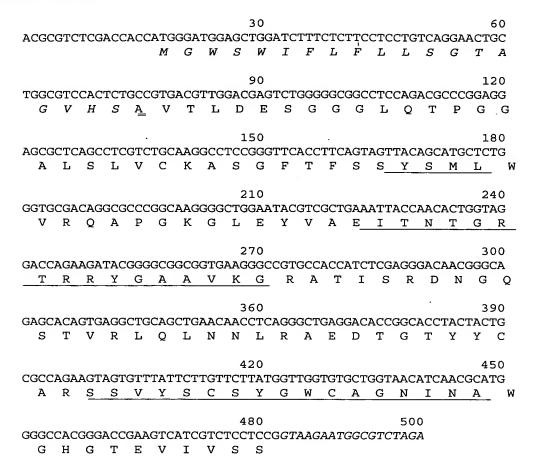
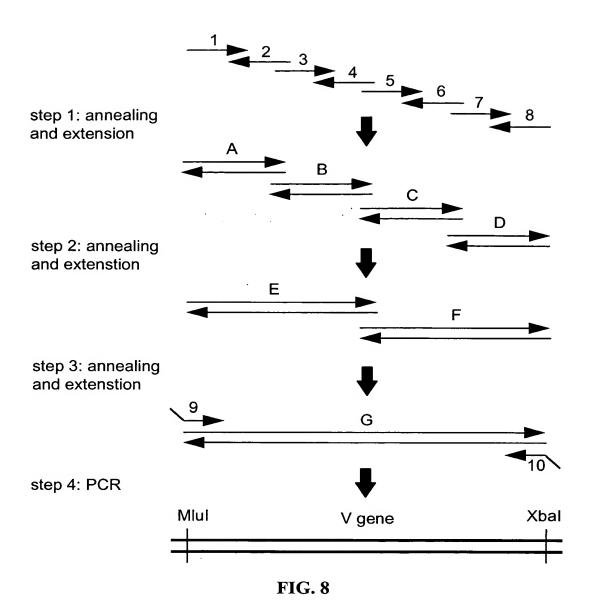


FIG. 7B



#### (A) For HuB1 Vλ

5'-GACTCGTCTAGAGGTTGTGAG-3'

```
Primer 1:
5'-CTAGCCACGCGTCCACCATGGAGAAAGACACTCCTGCTATGGGTCCTACTTCTCTGGGTTCCAGGTTC-3'
Primer 2:
5'-CCAGGGCCACTGACACTGAAGGCGGCTGAGTCAGCTCAGAGCTACCTGTGGAACCTGGAACCCAGAGAAG-3'
5'-CTTCAGTGTCAGTGGCCCTGGGACAGACCGTCAGGATCACCTGCTCCGGAGGTTACAGCGGCTATTATGGC-3'
5'-GTTGTCATAAATCACAGTGACAGGAGCCTGGCCAGGTTTCTGCTGGTACCAGCCATAATAGCCGCTGTAAC-3'
5'-CCTGTCACTGTGATTTATGACAACACCAGGAGACCCTCGGACATCCCTTCACGATTCTCCGGTTCCAAATCCG-3'
Primer 6:
\verb§5'-CCTCGTCCTCGGCTTGGACTCCAGTGATGGTTAATGTGGCTGTGGAGCCGGATTTGGAACCGGAGAATC-3'
5'-GAGTCCAAGCCGAGGACGAGCTGACTATTACTGTGGGACCTGGGACAGCAGCCGTGTTGGTATATTTGGAGG-3'
Primer 8:
5'-GACTCGTCTAGAGGGAGAGAGACTCACCTAGGACGGTCAGCTTTGTCCCACCTCCAAATATACCAACACGGC-3'
Primer 9:
5'-CTAGCCACGCGTCCACCATG-3'
Primer 10:
5'-GACTCGTCTAGAGGGAGAAG-3'
(B) For HuB1 VH
5'-CTAGCCACGCTCCACCATGGATGGAGCTGGATCTTTCTCTTCTCTCTGTCAGGAACTGCTGGCGTCCACTCTCAGG-3'
5'-GAGCCTGAGGCTTCCTCCAGGCTGCACGAGTCCACCACCACCTGCACCTGAGAGTGGACGCCAGCAG-3'
Primer 3:
\verb§5'-CCTGGAGGAAGCCTCAGGCTCAGCTCGCGCCTCCGGGTTCACCTTCAGTAGTTACAGCATGCTCTGGGTGCGACAGG-3'
5'-CTTCTGGTCCTACCAGTGTTGGTAATTTCAGCGACGTATTCCAGTCCCTTGCCAGGCGCCTGTCGCACCCAGAGCATG-3'
Primer 5
5'-CCAACACTGGTAGGACCAGAAGATACGGAGCTGCGGTGAAGGGCCGTGCCACCATCTCTAGGGACAACGCCAAGAACAC-3'
Primer 6:
\verb§5'-GGCGCAGTACTACACGGCGGTGTCCTCAGCCCTGAGGCTGTTCATCTGCAGGTACACTGTGTTCTTGGCGTTGTCCCTA-3'
Primer 8:
5'-GACTCGTCTAGAGGTTGTGAGGACTCACCGGAGGAGACGGTGACCAGGGTTCCCTGGCCCCATGCGTTGATGTTACCAG-3'
Primer 9:
5'-CTAGCCACGCGTCCACCATG-3'
Primer 10:
```

## (A) Vλ mini-exon of humanized B1

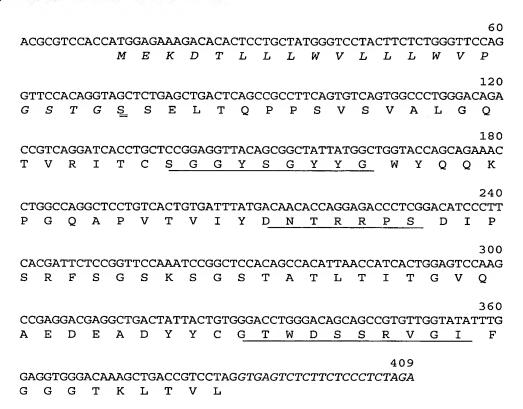
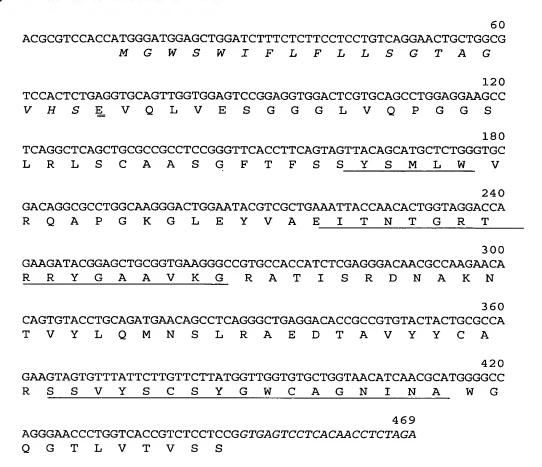


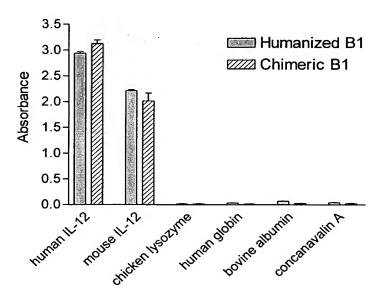
FIG. 10A

#### (B) VH mini-exon of humanized B1

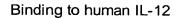


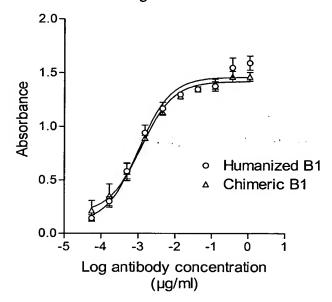
**FIG. 10B** 

## Binding to various proteins



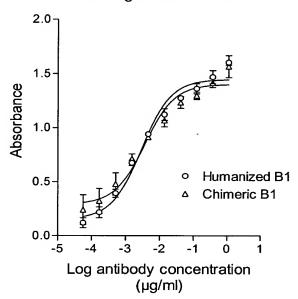
**FIG. 11** 





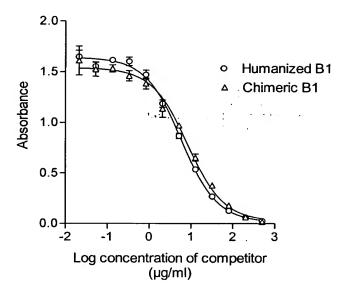
**(B)** 

## Binding to mouse IL-12



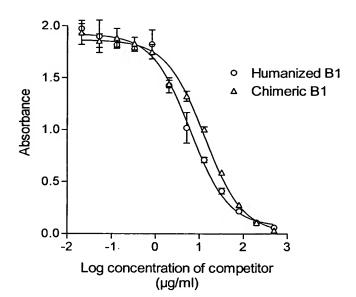
**FIG. 12** 

# Binding of biotinylated HuB1 to human IL-12



**(B)** 

## Binding of biotinylated HuB1 to mouse IL-12



**FIG. 13** 

## [DD2 Vλ mini-exon]

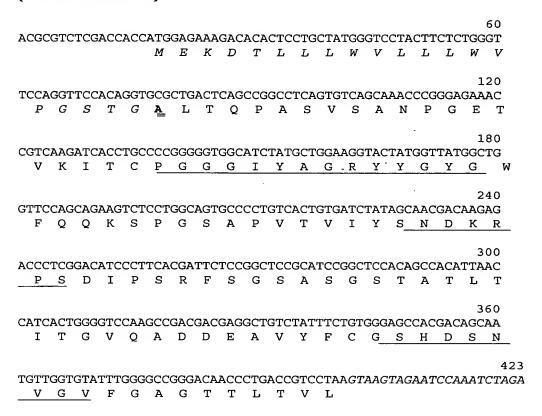


Fig. 14A

## [DD2 VH mini exon]

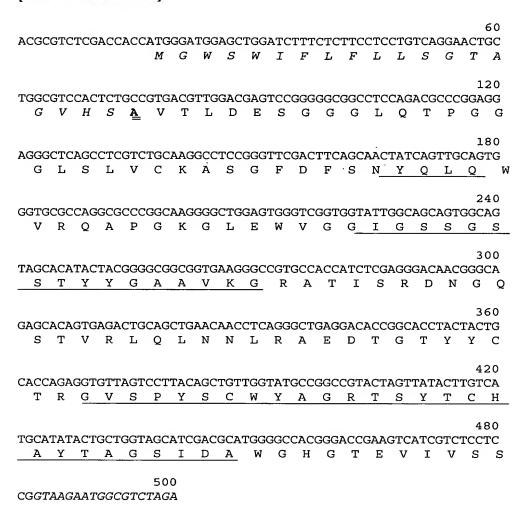


Fig. 14B

## (A) Vλ

	1	2	3	4
	123456789 012345678	9 0123456777789	01234567899	0123456789
		ABC	Α	
DD2	**ALTQPAS *VSANPGET	V KITCPGGGIYAGR	YYGYG WFQQKS	PGSAPVTVIY
HuDD2	SSELTQDPA *VSVALGQT	V RITCPGGGIYAGR	YYGYGWF QQK*	PGQAPVTVIY
DPL16	SSELTODPA *VSVALGOT			
	5 6	7 8	9	
	0123456789 01234567		<del>-</del>	2156700
DD2	SNDKRPSDIP SRFSGSAS			
HuDD2	SNDKRPSDIP SRFSGSAS	· -		
DPL16/Jλ2				
DPLITO/UNZ	GIP DRESGSSS	GN TASHILIGAQ A	EDEADIIC	FG
	1			
	0			
	01234567			
DD2	AGTTLTVL			
HuDD2	GGTKLTVL			
Јλ2	GGTKLTVL			
(B) VH				
	1	2 3	4	
	123456789 012345678	9 0123456789 01:	234567899 012	3456789
DD2	AVTLDESGG GLQTPGGGL			
HuDD2	EVQLVESGG GLVQPGGSL	_		_
DP-54	EVQLVESGG GLVQPGGSL			
0-				.022
	5 6	7 8	9	
	0123456789 01234567		-	0122456799
	0123430709 01234307	0) 0123430707 0.	ABC	0123430709
DD2	IGSSGSSTYY GAAVKGR	ידד פאראמטפייעים ז.		VVCTDCVCDV
HuDD2	IGSSGSSTYY GAAVKGRA			
DP-54	RF			
DI - J4	RF	II SKUMAKNOHI D	ZWM2HKWEDIKA	IICAR
	1			
	0			
	<u>-</u>	01004567000100		
	0000000000123456789	0123456/890123		
DD3	ABCDEFGHI	TD 1 1/01/02/2017	•	
DD2	SCWYAGRTSYTCHAYTAGS		į.	
HuDD2	SCWYAGRTSYTCHAYTAGS			
JH1		WGQGTLVTVSS		`

Fig. 15

#### (A) For HuDD2 Vλ

```
Primer 1
5'-ACGCGTCCACCATGGAGAAAGACACTCCTGCTGTGGGTCCTACTTCTCTGGGTTCCAGGTTCCACAGGTTC-3'
Primer 2
\verb§5'-CCTGACTGTCCCAAGGCCACAGACACAGGCTCCTGAGTCAGAAGAACCTGTGGAACCTGGAAC-3'
5'-CCTTGGGACAGACAGTCAGGATCACATGCCCCGGAGGTGGCATCTATGCTGGACGCTACTATGGTTATGCTG-3'
5'-CGTTGCTATAGATGACAGTTACAGGGGCCTGTCCTGGCTTCTGCTGGAACCATAACCATAGTAGCG-3'
Primer 5
5'-CTGTAACTGTCATCTATAGCAACGACAAGAGACCCTCGGACATCCCTTCACGATTCTCTGGCTCCGCATC-3'
Primer 6
\verb§5'-CATCTTCCGCCTGAGCCCCAGTGATGGTCAAGGAAGCTGTGGAGCCTGATGCGGAGCCAGAGAATCGTG-3'
5'-GGCTCAGGCGGAAGATGAGGCTGACTATTACTGTGGGAGCCACGACAGCAATGTTGGTGTATTTGG-3'
Primer 8
5'-TCTAGAGGGAGAAGACTCACCTAGGACGGTCAGCTTTGTCCCACCGCCAAATACACCAACATTGCTGTC-3'
Primer 9
5'-CTACGAACGCGTCCACCATGGAGAAAG-3'
5'-GACTTCTCTAGAGGGAGAAGAGACTCACC-3'
(B) For HuDD2 VH
5'-ACGCGTCCACCATGGGATGGAGCTGGATCTTTCTCTTCCTCCTGTCAGGAACTGCTGGCGTGCACTCTGAGGTGCAGCTG-3'
Primer 2
5'-GGCTGCACAGGAGAGTCTCAGGGACCCCCCAGGCTGGACCAAGCCTCCCCCAGACTCCACCAGCTGCACCTCAGAGTGCA-3'
Primer 3
\verb§5'-TGAGACTCTCCTGTGCAGCCTCTGGATTCGACTTTAGTAACTATCAGTTGCAGTGGGTCCGCCAGGCTCCAGGGAAGGGG-3'
Primer 5
5'-CATACTACGGAGCTGCGGTTAAGGGCCGAGCCACCATCTCCAGAGACAACGCCAAGAACTCAGTGTATCTGCAAATGAAC-3'
Primer 6
\verb§5'-CTGTAAGGACTAACACCTCTGGTACAGTAATACACAGCCGTGTCCTCGGCTCTCAGGCTGTTCATTTGCAGATACACTGA-3'
5'-AGAGGTGTTAGTCCTTACAGCTGTTGGTATGCCGGCCGTACTAGTTATACTTGTCATGCATATACTGCTGGTAGCATCGA-3'
\verb§5'-TCTAGAAGTACAGCAGACTCACCTGAGGAGACGGTGACCAGGGTTCCCTGGCCCCATGCGTCGATGCTACCAGCAGTATA-3'
Primer 9
5'-CTACGAACGCGTCCACCATGGGATGG-3'
Primer 10
```

Fig. 16

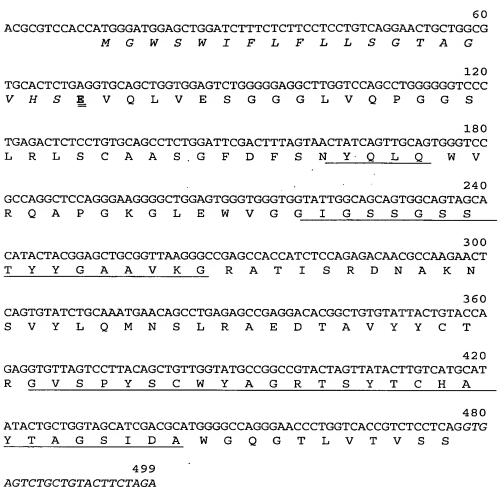
5'-GACTTCTCTAGAAGTACAGCAGACTCAC-3'

## [HuDD2 Vλ mini exon]

	~~~	maa		3 m.a	a.a		a	3 G 3	ama	ama	CITICO	maa	ото	C/III A	omm.	ama	maa	amm	60 CCAG
AC	GCG	TCC	ACC.						-	-		-	GIC V					GTT.	
				M	$\boldsymbol{E}$	K	D	T	$\boldsymbol{L}$	L	L	W	V	L	$\boldsymbol{L}$	$oldsymbol{L}$	W	V	P
																			100
<b>о</b> т	maa	202	aam	mam	m~m	~ ~ ~	ОШО	3 OM	~ ~ ~	C 2 C	COT		ОШО	mam	ama	aaa	mma	<b>~~</b>	120
																			CAGA
G	s	T	G	<u>s</u>	S	E	L	Т	Q	D	P	Α	V	S	V	Α	L	G	Q
																			180
CA	GTC.	AGG.	ATC.	ACA	TGC	CCC	GGA	GGT	GGC	ATC	TAT	'GCT	GGA	CGC	TAC	TAT	GGT	TAT	GGCT
Т	V	R	Ι	$\mathbf{T}$	С	<u>P</u> _	G	G	G	I	Y	A	G	R	Y	Y	G	Y	<u>G</u>
														٠.					
														-					240
GG	TTC	CAG	CAG	AAG	CCA	GGA	CAG	GCC	CCT	GTA	ACT	GTC	ATC	TAT.	AGC.	AAC	GAC.	AAG	AGAC
W	F	Q	Q	K	P	G	0	Α	P	v	Т	V	Ι	Y	s	N	D	K	R
																			300
CC	TCG	GAC	ATC	CCT	TCA	CGA	TTC	TCT	GGC	TCC	GCA	TCA	GGC	TCC	ACA	GCT	TCC	тта	ACCA
P P		D	T	P	s	R	F	s	G	s	A	s	G	S	т	A	S	L	Т
	<del>-</del>	_	_	-	_		-	_	•	~	••	_	Ŭ	_	•	••	_	_	-
																			360
тС	л ("Τ"	aaa	o Omo	רא כי	aca.	מאים	CAT	CAC	C CT	CAC	ጥለጥ	ጥ አ 🕶	ጥርጥ	ccc	אככ	CAC	~	7 C C	AATG
								GAG E		_			161						
I	Т	G	Α	Q	A	E	D	E	A	D	Y	Y	C	G_	S	H	D	s	<u>N</u>
																			421
	TTGGTGTATTTGGCGGTGGGACAAAGCTGACCGTCCTAG <i>GTGAGTCTCTTCTCCCTCTAGA</i>																		
<u>V</u>	_G_	<u>V</u>	F	G	G	G	Т	K	L	$\mathbf{T}$	V	L							

Fig. 17A

#### [HuDD2 VH mini exon]



**AGTCTGCTGTACTTCTAGA** 

**Fig. 17B** 

## Binding to various proteins

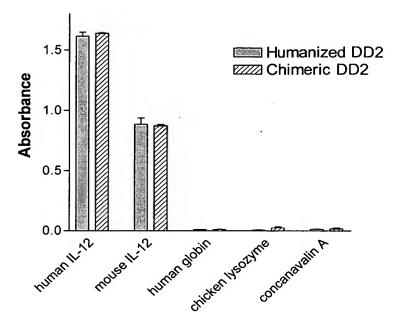
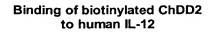
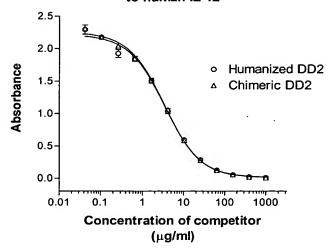


Fig. 18

**(A)** 





**(B)** 

## Binding of biotinylated ChDD2 to mouse IL12

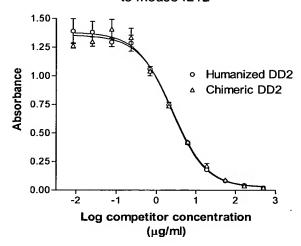
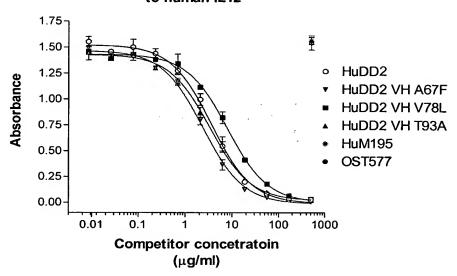


Fig. 19

**(A)** 





**(B)** 

## Binding of biotinylated ChDD2 to human IL12

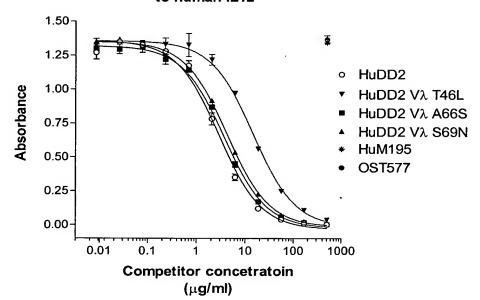


Fig. 20

## [D3 V\(\lambda\) mini-exon]

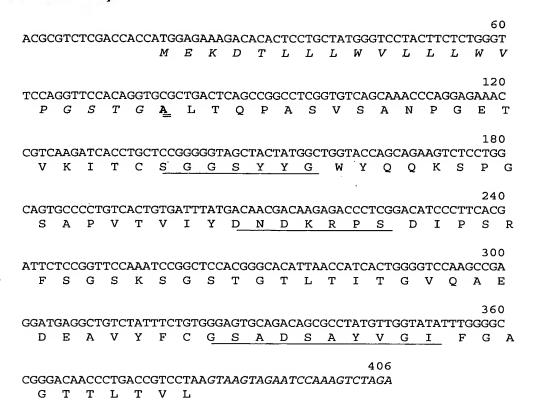


Fig. 21A

#### [D3 VH mini-exon]

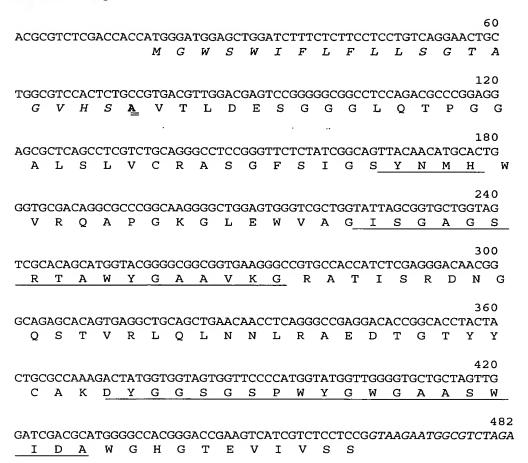


Fig. 21B

```
[V\]
                   123456789 0123456789 0123456789 01234567899 0123456789
                   **ALTQPAS *VSANPGETV KITCSGGS** **YYGWYQQKS PGSAPVTVIY
SSELTQDPA *VSVALGQTV RITCSGGS** **YYGWYQQK* PGQAPVT_VIY
Chicken D3
Humanized D3
3-230IIIB237
                   SSELTQDPA *VSVALGQTV RITC----- -- WYQQK* PGQAPVLVIY
                                                          8
                    0123456789 0123456789 0123456789 0123456789 01234556789
                   DNDKRPSDIP SRFSGSKSGS TGTLTITGVQ AEDEAVYFCG SADSAYVGIFG
Chicken D3
Humanized D3 DNDKRPSDIP SRFSGSKSGS TGSLTITGAQ ABDEADYYCG SADSAYVGIFG 3-230IIIB237/J\(\text{J}\)2 ------GIP DRFSGSSSGN TASLTITGAQ AEDEADYYC- ------FG
                    O
                   01234567
Chicken D3
                   AGTTLTVL
Humanized D3
                   GGTKLTVL
J<sub>2</sub>
                   GGTKLTVL
[VH]
                   123456789 0123456789 0123456789 0123456789 0123456789
Chicken D3
                   AVTLDESGG GLQTPGGALS LVCRASGFSI GSYNMH WVRQ APGKGLEWVA
                   EVQLLESGG GLVQPGGSLR LSCAASGFSI GSYNMHWVRQ APGKGLEWVA EVQLLESGG GLVQPGGSLR LSCAASGFTF S----WVRQ APGKGLEWVS
Humanized D3
ha316
                                                            8
                   012223456789 0123456789 0123456789 0122223456789 0123456789
                                                               ABC
Chicken D3
                   GISGAGSRTAWY GAAVKG RATI SRDNGQSTVR LQLNNLRAEDTGT YYCAKDYGGS
Humanized D3
                   GISGAGSRTAWY GAAVKGRA TI SRDNAKNTVY LOMNSLRAEDTAV YYCAKDYGGS
                   ----- ----- SRDNSKNTLY LQMNSLRAEDTAV YYCAK----
ha316
                                              7
                   000000000000123456789 0123
                    ABCDEFGHIJKL
Chicken D3
                   GSPWYGWGAASWIDA WGHGTEV IVSS
Humanized D3
                   GSPWYGWGAASWIDAWGQGTLV TVSS
ha316
                    -----WGQGTLV TVSS
```

**Fig. 22** 

## [HuD3 Vλ mini-exon]

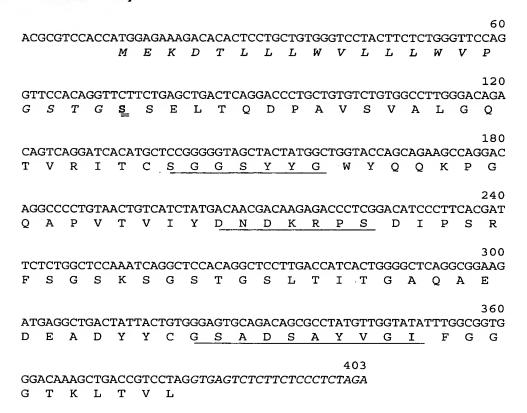
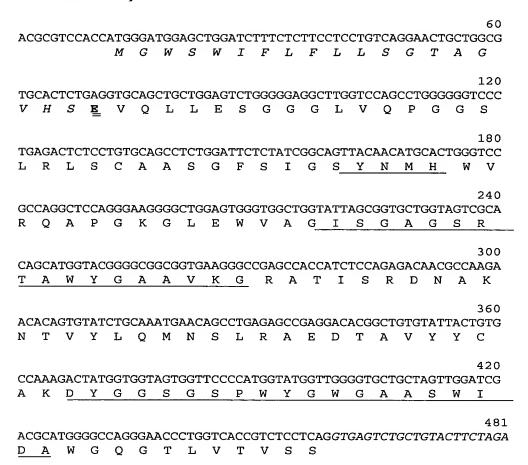


Fig. 23A

## [HuD3 VH mini-exon]



**Fig. 23B** 

## Binding to L-selectin

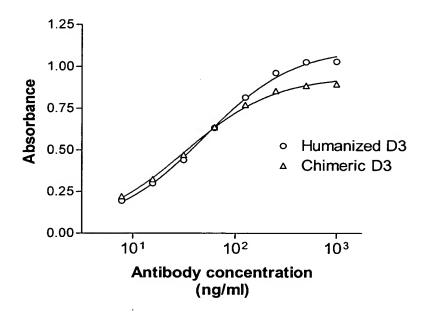


Fig. 24

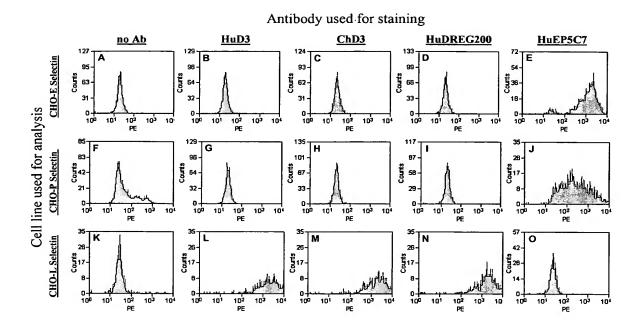
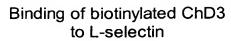
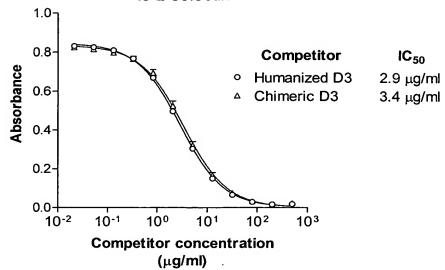


Fig. 25





**(B)** 

# Binding of biotinylated ChD3 to L-selectin

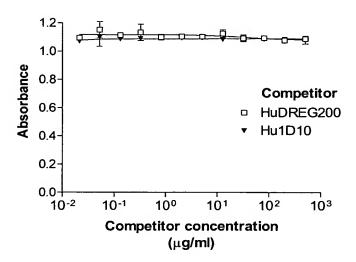


Fig. 26